

region of **adenovirus 5**. This construct directed high levels of synthesis of **p53** in HeLa cells.

L9 ANSWER 42 OF 42 MEDLINE on STN DUPLICATE 15
ACCESSION NUMBER: 95134785 MEDLINE
DOCUMENT NUMBER: PubMed ID: 7833367
TITLE: Development and characterization of recombinant **adenoviruses** encoding human **p53** for **gene therapy** of cancer.
AUTHOR: Wills K N; Maneval D C; Menzel P; Harris M P; Sutjipto S; Vaillancourt M T; Huang W M; Johnson D E; Anderson S C; Wen S F; +
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AB We have constructed recombinant human **adenoviruses** that express wild-type human **p53** under the control of either the Ad 2 major late promoter (MLP) or the human cytomegalovirus (CMV) immediate early gene promoter. Each construct replaces the Ad 5 Ela and Elb coding sequences necessary for viral replication with the **p53** cDNA and MLP or CMV promoter. These **p53**/Ad recombinants are able to express **p53** protein in a dose-dependent manner in infected human cancer cells. Tumor suppressor activity of the expressed **p53** protein was assayed by several methods. [3H]Thymidine incorporation assays showed that the recombinant **adenoviruses** were capable of inhibiting DNA synthesis in a **p53**-specific, dose-dependent fashion. Ex vivo treatment of Saos-2 tumor cells, followed by injection of the treated cells into nude mice, led to complete tumor suppression using the MLP/**p53** recombinant. Following a single injection of CMV/**p53** recombinant **adenovirus** into the peritumoral space surrounding an in vivo established tumor derived from a human small cell lung carcinoma cell line (NIH-H69), we were able to detect **p53** mRNA in the tumors at 2 and 7 days post-injection. Continued treatment of established H69 tumors with MLP/**p53** recombinant led to reduced tumor growth and increased survival time compared to control treated animals. These results indicate that recombinant **adenoviruses** expressing wild-type **p53** may be useful vectors for **gene therapy** of human cancer.

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FILE 'MEDLINE' ENTERED AT 06:24:50 ON 28 SEP 2004

L1 29300 S ADENOVIR?
L2 681 S L1 AND (CMV OR CMV?)
L3 33 S L2 AND P53
L4 22 S L3 AND CANCER
L5 24 S L3 AND NEOPLASM
L6 14 S L5 AND TREATMENT
L7 19 S L5 AND GENE(W) THERAPY

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